

Figure 1. DNA Sequence of the human IL-1A gene. (GenBank Accession No. X03833)

-1437 AAGCTTCTAC CCTAGTCTGG TGCTACACTT ACATTGCTTA CATCCAAGTG TGGTTATTTT
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 -717 TGGTCCTTGG TAGAGGGCTA CTTTACTGTA ACAGGGCCAG GGTGGAGAGT TCTCTCCTGA
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 477 GGCTCATTTT CCCTCAAAAG TTGCCAGGAG CTGCCAAGTA TTCTGCCAAT TCACCCTGGA
 417 GCACAATCAA CAAATTCAGC CAGAACACAA CTACAGCTAC TATTAGAACT ATTATTATTA
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 297 TAGAACTTG ATAAGTTTCC CGCGCTTCCC TTTTCTAAG ACTACATGTT TGTATCTTA
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 177 AATAATATCA GCTATGCCAT CTTTCACTAT TTTAGCCAGT ATCGAGTTGA ATGAACATAG
 117 AAAAATACAA AACTGAATTC TTCCCTGTAA ATTCCCGT TTAGCAGCGC ACTGTAGCC
 57 ACGTAGCCAC GCCTACTTAA GACAATTACA AAAGGCGAAG AAGACTGACT CAGGCTTAAG
 4 CTGCCAGCCA GAGAGGGAGT CATTTCATG GCGTTTGAGT CAGCAAAGGT ATTGTCCTCA
 64 CATCTCTGGC TATTAAAGTA TTTTCTGTTG TTGTTTTTCT CTTTGGCTGT TTTCTCTCAC
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 244 GATTGCTTTT ACTGAGGGAC GGCAGAACTA GTTTCCTATG AGGGCATGGG TGAATACAAC
 304 TGAGGCTTCT CATGGGAGGG AATCTCTACT ATCCAAAATT ATTAGGAGAA AATTGAAAAT
 364 TTCCAACCTCT GTCTCTCTCT TACCTCTGTG TAAGGCAAAT ACCTTATTCT TGTGGTGTGTT
 424 TTGTAAACCTC TTCAAACCTT CATTGATTGA ATGCCTGTTT TGGCAATACA TTAGGTGTTG
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 544 AGACAGGTAC ATAGCAAAC AATTCAAAGG AGCTAGAAGA TGGAGAAAAT GCTGAATGTG
 604 GACTAAGTCA TTCAACAAAG TTTTCAGGAA GCACAAAGAG GAGGGGCTCC CCTCACAGAT
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 1684 ACATGAAATA ACAACAATTA CATTCTCATC ATCTTATTTT GACAGTGAAA ATGAAGAAGA
 1744 CAGTTCCTCC ATTGATCATC TGTCTCTGAA TCAGGTAAGC AAATGACTGT AATTCTCATG
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 1924 ACCAGCTGCC TCTCTAAACT AATAGTTGAT GTGCATTGGC TTCTCCCAGA ACAGAGCAGA
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 2044 ATGCCATCCT GAGAAAGCCC CGCAGGCCGC TTCACCAGGT GTCTGTCTCC TAACGTGATG
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 7504 GGGTAGGACA TTCAGAGGAG GGGGCGGGTC GTGGTTGTGG GTTTTTGGGT AGGACATTCA
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 7804 ATATGCAGAG ACTAGTGCTT GCAGAGCTTG CATTTGGATT TCATTTGAGG TACAATGAAA
 7864 ACCCATTAAT GGGTTTCACA CAGTGCAATG GCCTGACCTC ACTTATATTT CCTAAAATAG

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|-------|------------|-------------|------------|------------|------------|------------|
| 7924 | AAAACAGATC | AGAAGGAAGG | CAATAGAGAA | GCAGAAAGTC | CAATGAGGAG | GTTTCACAGC |
| 7984 | AGTCATGGGG | GTGGGGTAAG | GAAAAGAAGT | GGAAAGAAAC | AGACAGAATT | GGGTATATTT |
| 8044 | TTGGAGATAG | AACCAACAGA | AGGAAGAGGA | GAAACAACAT | TTACTGAGAA | GGGAAAAAGT |
| 8104 | AGGAGAGGAA | TAGGTTTGGG | AAATAAATCC | TGCTGACATT | GGAAACCCCA | AGGAAGCCTC |
| 8164 | AAAAGTATAT | TTACTTGCTT | TAGATTTAAA | AGAATAGGAA | AGAAGCATCT | CAACTTGGAA |
| 8224 | TTTGAAATCT | ATTTTTCCAT | AAAAGTATTG | TTAAATTCTA | CTCATACTCA | CAAGAAAAGT |
| 8284 | ACATTCTAAA | GAGTATATTG | AAAGAGTTTA | CTGATATACT | TAGGAATTTT | GTGTGTATGT |
| 8344 | GTGTGTGTGT | ATGTGTGTGT | GTGTGTTTAA | CCTTCAATTG | TTGACTTAAA | TACTGAGATA |
| 8404 | AATGTCATCT | AAATGCTAAA | TTGATTTCCC | AAAGGTATGA | TTTGTTCACT | TGGAGATCAA |
| 8464 | AATGTTTAGG | GGGCTTAGAA | TCACTGTAGT | GCTCAGATT | GATGCAAAAT | GTCTTAGGCC |
| 8524 | TATGTTGAAG | GCAGGACAGA | AACAATGTTT | CCCTCCTACC | TGCCTGGATA | CAGTAAGATA |
| 8584 | CTAGTGTAC | TGACAATCTT | CATAACTAAT | TTAGATCTCT | CTCCAATCAA | CTAAGGAAAT |
| 8644 | CAACTCTTAT | TAATAGACTG | GGCCACACAT | CTACTAGGCA | TGTAATAAAT | GCTTGCTGAA |
| 8704 | TGAACAAATG | AATGAAGAGC | CTATAGCATC | ATGTTACAGC | CATAGTCCTA | AAGTGGTGT |
| 8764 | TCTCATGAAG | GCCAAATGCT | AAGGGATTGA | GCTTCAGTCC | TTTTTCTAAC | ATCTTGTCT |
| 8824 | CTAACAGAAT | TCTCTTCTTT | TCTTCATAGG | AGATGCCTGA | GATACCCAAA | ACCATCACAG |
| 8884 | GTAGTGAGAC | CAACCTCCTC | TTCTTCTGGG | AAACTCACGG | CACTAAGAAC | TATTTACAT |
| 8944 | CAGTTGCCCA | TCCAACTTG | TTTATTGCCA | CAAAGCAAGA | CTACTGGGTG | TGCTTGGCAG |
| 9004 | GGGGGCCACC | CTCTATCACT | GACTTTCAGA | TACTGGAAAA | CCAGGCGTAG | GTCTGGAGTC |
| 9064 | TCACTTGTCT | CACTTGTGCA | GTGTTGACAG | TTCATATGTA | CCATGTACAT | GAAGAAGCTA |
| 9124 | AATCCTTTAC | TGTTAGTCAT | TTGCTGAGCA | TGTACTGAGC | CTTGTAATTC | TAAATGAATG |
| 9184 | TTTACACTCT | TTGTAAGAGT | GGAACCAACA | CTAACATATA | ATGTTGTTAT | TTAAGAACA |
| 9244 | CCCTATATTT | TGCATAGTAC | CAATCATTTT | AATTATTATT | CTTCATAACA | ATTTTAGGAG |
| 9304 | GACCAGAGCT | ACTGACTATG | GCTACCAAAA | AGACTCTACC | CATATTACAG | ATGGGCAAA |
| 9364 | TAAGGCATAA | GAAAACCTAAG | AAATATGCAC | AATAGCAGTT | GAAACAAGAA | GCCACAGACC |
| 9424 | TAGGATTTCA | TGATTTTCAAT | TCAACTGTTT | GCCTTCTGCT | TTTAAGTTGC | TGATGAATC |
| 9484 | TTAATCAAAT | AGCATAAGTT | TCTGGGACCT | CAGTTTATC | ATTTTCAAAA | TGGAGGGAAT |
| 9544 | AATACCTAAG | CCTTCCTGCC | GCAACAGTTT | TTTATGCTAA | TCAGGGAGGT | CATTTTGGTA |
| 9604 | AAATACTTCT | CGAAGCCGAG | CCTCAAGATG | AAGGCAAAGC | ACGAAATGTT | ATTTTTTAAT |
| 9664 | TATTATTTAT | ATATGTATTT | ATAAATATAT | TTAAGATAAT | TATAATATAC | TATATTTATG |
| 9724 | GGAACCCCTT | CATCCTCTGA | GTGTGACCAG | GCATCCTCCA | CAATAGCAGA | CAGTGTTTTT |
| 9784 | TGGGATAAGT | AAGTTTGATT | TCATTAATAC | AGGGCATTTT | GGTCCAAGTT | GTGCTTATCC |
| 9844 | CATAGCCAGG | AAACTCTGCA | TTCTAGTACT | TGGGAGACCT | GTAATCATAT | AATAAATGTA |
| 9904 | CATTAATTAC | CTTGAGCCAG | TAATTGGTCC | GATCTTTGAC | TCTTTTGCCA | TTAACTTAC |
| 9964 | CTGGGCATTC | TTGTTTCATT | CAATTCCACC | TGCAATCAAG | TCCTACAAGC | TAAAATTAGA |
| 10024 | TGAACCTAAC | TTTGACAACC | ATGAGACCAC | TGTTATCAAA | ACTTTCTTTT | CTGGAATGTA |
| 10084 | ATCAATGTTT | CTTCTAGGTT | CTAAAAATTG | TGATCAGACC | ATAATGTTAC | ATTATTATCA |
| 10144 | ACAATAGTGA | TTGATAGAGT | GTTATCAGTC | ATAACTAAAT | AAAGCTTGCA | ACAAAATTCT |
| 10204 | CTGACACATA | GTTATTCATT | GCCTTAATCA | TTATTTTACT | GCATGGTAAT | TAGGGACAAA |
| 10264 | TGGTAAATGT | TTACATAAAT | AATTGTATTT | AGTGTTACTT | TATAAAATCA | AACCAAGATT |
| 10324 | TTATATTTTT | TTCTCCTCTT | TGTTAGCTGC | CAGTATGCAT | AAATGGCATT | AAGAATGATA |
| 10384 | ATATTTCCGG | GTTCACTTAA | AGCTCATATT | ACACATACAC | AAAACATGTG | TTCCCATCTT |
| 10444 | TATACAAACT | CACACATACA | GAGCTACATT | AAAAACAAC | AATAGGCCAG | GCACGGTGGC |
| 10504 | TCAGACCTGT | AATCCCAGCA | CTTTGGGAGG | | | |

Figure 2. DNA Sequence of the human IL-1B gene. (GenBank Accession No. X04500)

-1933 AGAAAGAAAG AGAGAGAGAA AGAAAAGAAA GAGGAAGGAA GGAAGGAAGG AAGAAAGACA
 -1873 GGCTCTGAGG AAGGTGGCAG TTCCTACAAC GGGAGAACCA GTGGTTAATT TGCAAAGTGG
 -1813 ATCCTGTGGA GGCANNCAGA GGAGTCCCCT AGGCCACCCA GACAGGGCTT TTAGCTATCT
 -1753 GCAGGCCAGA CACCAAATTT CAGGAGGGCT CAGTGTTAGG AATGGATTAT GGCTTATCAA
 -1693 ATTCACAGGA AACTAACATG TTGAACAGCT TTTAGATTTT CTGTGGAAAA TATAACTTAC
 -1633 TAAAGATGGA GTTCTTGTGA CTGACTCCTG ATATCAAGAT ACTGGGAGCC AAATTAAAAA
 -1573 TCAGAAGGCT GCTTGGAGAG CAAGTCCATG AAATGCTCTT TTTCCACAG TAGAACCTAT
 -1513 TTCCCTCGTG TCTCAAATAC TTGCACAGAG GCTCACTCCC TTGGATAATG CAGAGCGAGC
 -1453 ACGATACCTG GCACATACTA ATTTGAATAA AATGCTGTCA AATTTCCATT CACCCATTCA
 -1393 AGCAGCAAAC TCTATCTCAC CTGAATGTAC ATGCCAGGCA CTGTGCTAGA CTGGCTCAA
 -1333 AAAGATTTCA GTTTCCTGGA GGAACCAGGA GGGCAAGGTT TCAACTCAGT GCTATAAGAA
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 -1213 GGGCAGATCA CAAGGTCAGG AGATCGAGAC CATCCTGGCT AACATGGTGA AACCTGTCT
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 -1093 TGGGGAGGCT GAGGCAGGAG AATGGTGTGA ACCCGGGAGG CGGAAC TTGC AGGGGGCCGA
 -1033 GATCGTGCCA CTGCACTCCA GCCTGGGCGA CAGAGTGAGA CTCTGTCTCA AAAAAAAAAA
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 -913 ACTGTTTATA ACCTGGACTC TCATTCATTC TACAAATGGA GGGCTCCCCCT GGGCAGATCC
 -853 CTGGAGCAGG CACTTGTCTG GTGTCTCGGT TAAAGAGAAA CTGATAACTC TTGGTATTAC
 -793 CAAGAGATAG AGTCTCAGAT GGATATTCTT ACAGAAACAA TATTTCCACT TTTCAGAGTT
 -733 CACCAAAAAA TCATTTTAGG CAGAGCTCAT CTGGCATTGA TCTGGTTCAT CCATGAGATT
 -673 GGCTAGGGTA ACAGCACCTG GTCTTGCAAG GTTGTGTGAG CTTATCTCCA GGGTTGCCCC
 -613 AACTCCGTCA GGAGCCTGAA CCCTGCATAC CGTATGTTCT CTGCCCCAGC CAAGAAAGGT
 -553 CAATTTTCTC CTCAGAGGCT CCTGCAATTG ACAGAGAGCT CCCGAGGCAG AGAACAGCAC
 -493 CCAAGGTAGA GACCCACACC CTCAATACAG ACAGGGAGGG CTATTGGCCC TTCTATTGTAC
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Figure 3. DNA Sequence of the human IL-1RN gene. (GenBank Accession No. X64532)

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 793 GAAATCATGT TGGGGTGACT TCTCCCCAC TCTGACCTTT ATGTTTGTCT GGGCCGAGGC
 853 TGCAAGTCGG GCTCTGTGGG TGTATGAGTG ACAAGTCTCT CCCTTCCAGA TATGGGGACT
 913 GTCTGCTTCC CTAGGTTGCC TCTCCCTGCT CTGATCAGCT AGAAGCTCCA GGAGATCCTC
 973 CTGGAGGCCC CAGCAGGTGA TGTTTATCCC TCCAGACTGA GGCTAAATCT AGAAACTAGG
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 1693 GTGCCTTCCC TGGGAATCTC AGATGGGAAG CAAGTAAGGA GGGGAGTCAA ATGTGGGCTC
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 1813 CATCCATGCA TGCCGTGGGT ATACTAAAAT ACTATACCCC TGGAAGAGCT GGATGCAAAAT
 1873 TTGACAAGTT CTGGGGGACA CAGGAAGGTG CCAAGCACAA GGCTGGGCAC ATGGTGGCTG
 1933 TGCACTACAG CTGAGTCCTT TTCCTTTTCA GAATCTGGGA TGTTAACCAG AAGACCTTCT
 1993 ATCTGAGGAA CAACCAACTA GTTGTGGAT ACTTGCAAGG ACCAAATGTC AATTTAGAAG
 2053 GTGAGTGGTT GCCAGGAAAG CCAATGTATC TGGGCATCAC GTCACCTTGC CCGTCTGTCT
 2113 GCAGCAGCAT GGCCTGCCTG CACAAACCCT AGGTGCAATG TCCTAATCCT TGTTGGGTCT
 2173 TTGTATTCAA GTTTGAAGCT GGGAGGGCCT GGCTACTGAA GGGCACATAT GAGGGTAGCC
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 2773 TTTGTTGCTA CATGAGAGCA TGGAGGCCTC TTAGGGAGAG AGGAGGTTCA GAGACTCCTA
 2833 GGCTCCTGGT GGAGCCCCAC TCATGGCCTT GTTCATTTTC CCTGCCCTC AGCAACACTC
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 2953 ACAACATCCA GGAGACTCAG GCCTCTAGGA GTAAGTGGGT ACTCAGGCCT CTAGGAGTAA
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 3493 TGTGCCTGTC CTGTGTCAAG TCTGGTGATG AGACCAGACT CCAGCTGGAG GTAAAAACAT
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 3673 GGGGTCACTT TGGAAGCTGC ATTCAGCAGA GTGCCAGGCT TGCCTGGGC ATCCAAGGTG
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 4033 TTGCTCAAGG GCCCCCTAC CCACGCAGAC CTGCTGTCTT CTAGCAAAGC CCATCCTCAG
 4093 GACCTTTCTC TTCCAATCCT TATTGACTCA AATTGATTAG TTGGTGCTCC ACCCAGAGCC
 4153 CTGTGCTCCT TTATCTCATG TAATGTTAAT GGGTTTCCCA GCCCTGGGAA AACATGGCTT
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 5053 ACTGACCTGA GCGAGAACAG AAAGCAGGAC AAGCGCTTCG CCTTCATCCG CTCAGACAGT
 5113 GGCCCCACCA CCAGTTTGA GTCTGCCGCC TGCCCCGGTT GGTTCCTCTG CACAGCGATG
 5173 GAAGCTGACC AGCCCCGTCAG CCTCACCAAT ATGCCTGACG AAGGCGTCAT GGTCACCAAA
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 6133 GAATAAATTG CTCCTTGACA TTGTAGAGCT TCTGGCACTT GGAGACTGT ATGAAAGATG
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 6433 TTTGGGTATA GAGTGCTGAG GAAACTGAAA GACCAATGTG TYTTYCTTAC CCCAGAGGCT
 6493 GGCCTGCTG CTCTTCTCTG AGAGTTCTTT TCTTCCTTCA GCCTCACTCT CCCTGGATAA

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6553 CATGAGAGCA AATCTCTCTG CGGGG

09578534.052400

Figure 4

Genes for IL-1 α , IL-1 β , and IL-1 receptor antagonist are found in a cluster on chromosome 2

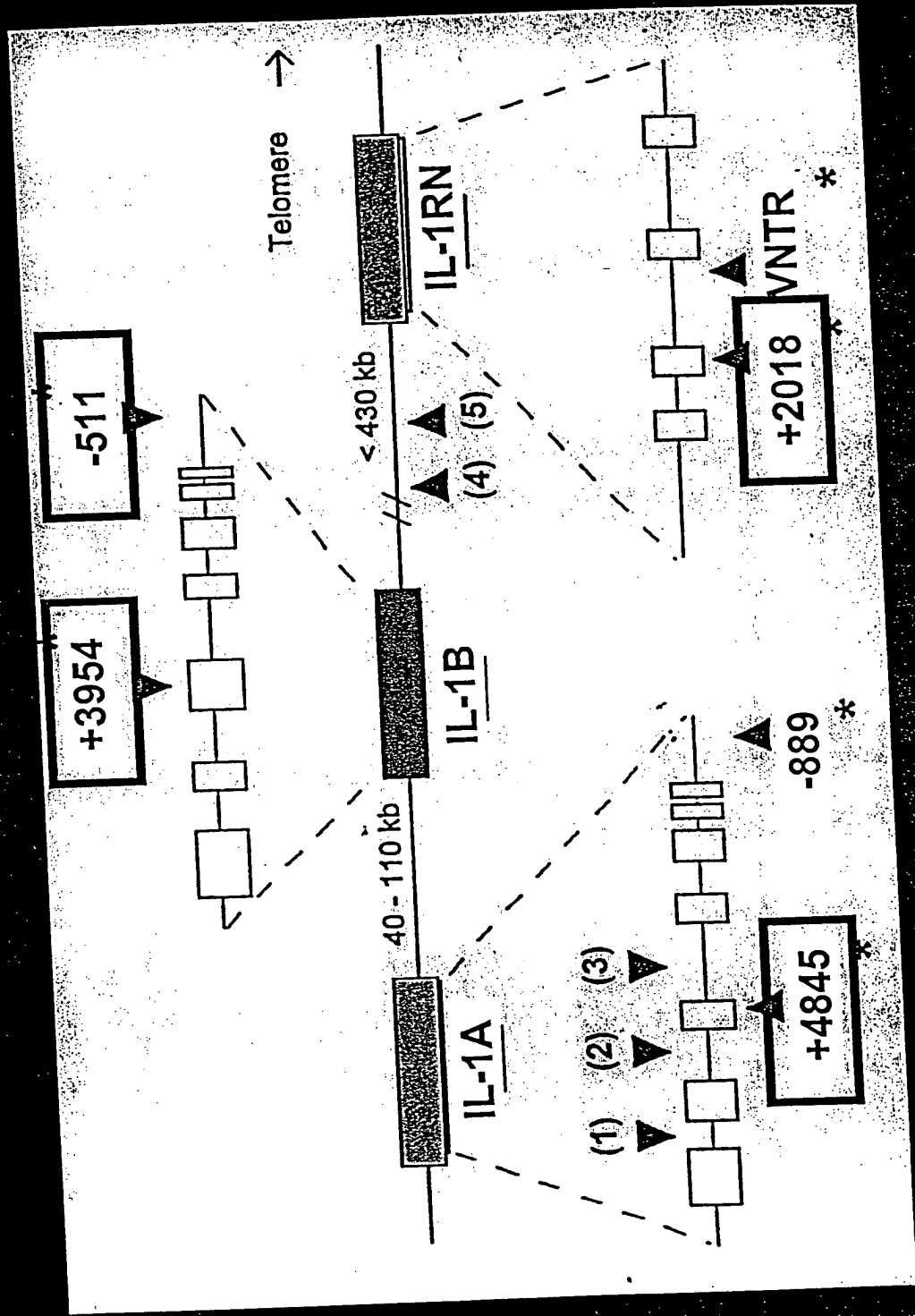


Figure 5

Disequilibrium Values within the IL-1 Gene Cluster

| | A(+4845) | B(+3954) | B(-511) | RN(+2018) |
|-----------|----------|----------|---------|-----------|
| A(+4845) | ---- | 0.804 | -0.264 | -0.207 |
| B(+3954) | 0.804 | ---- | -0.617 | -0.439 |
| B(-511) | -0.264 | -0.617 | ---- | 0.448 |
| RN(+2018) | -0.207 | -0.434 | 0.448 | ---- |

212 unrelated healthy Caucasians
Adapted from Cox et al. *Am J Human Genetics* 1998

Figure 6

Frequencies of Haplotype Patterns

IL-1A IL-1B IL-1RN
+4845 +3954 -511 +2018

1 1 1 1

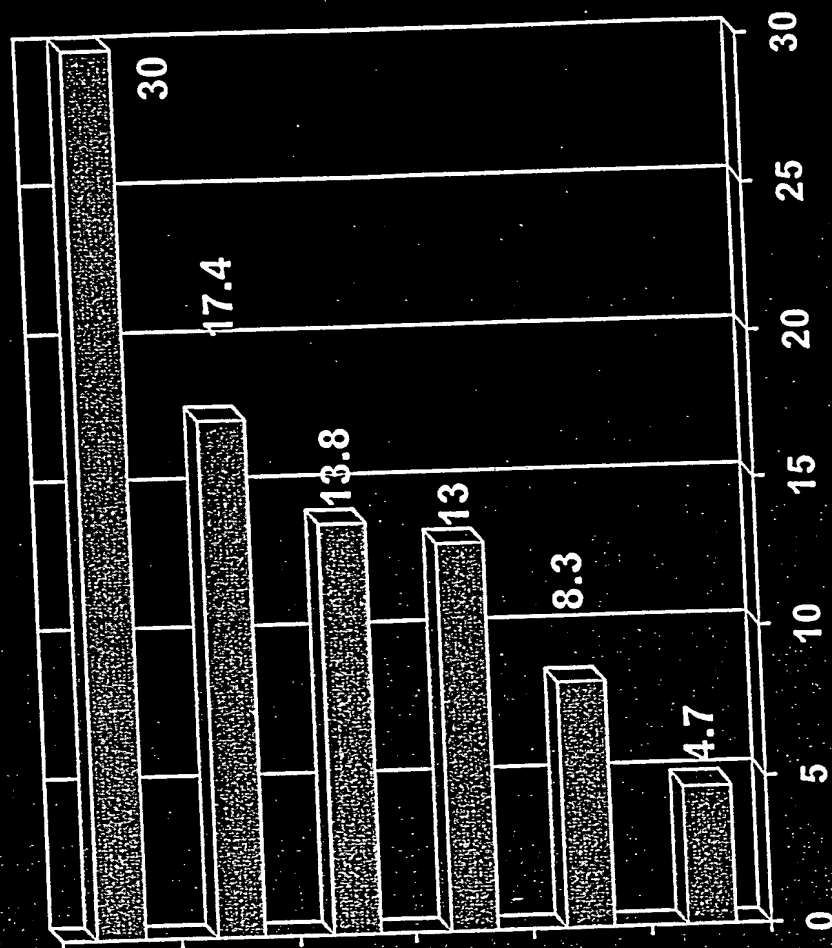
2 2 1 1

1 1 2 1

1 1 2 2

1 1 1 2

2 1 1 1



N=1343 Caucasians
Haplotype frequencies calculated
using Arlequin v. 1.1

Figure 7

A Different Pattern is Associated with Risk for Restenosis: Preliminary data

Kastrati, Schomig, et al 1999

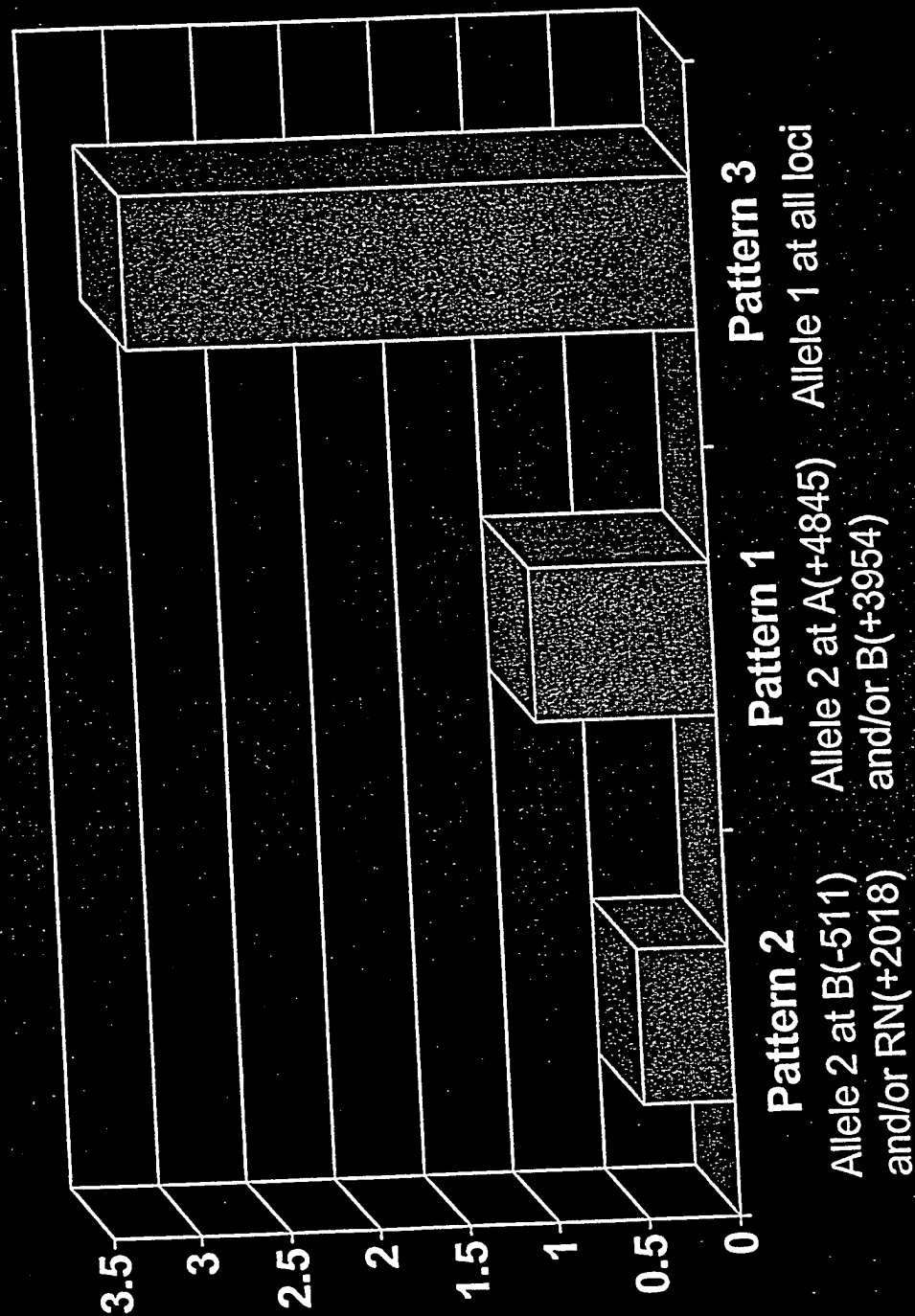
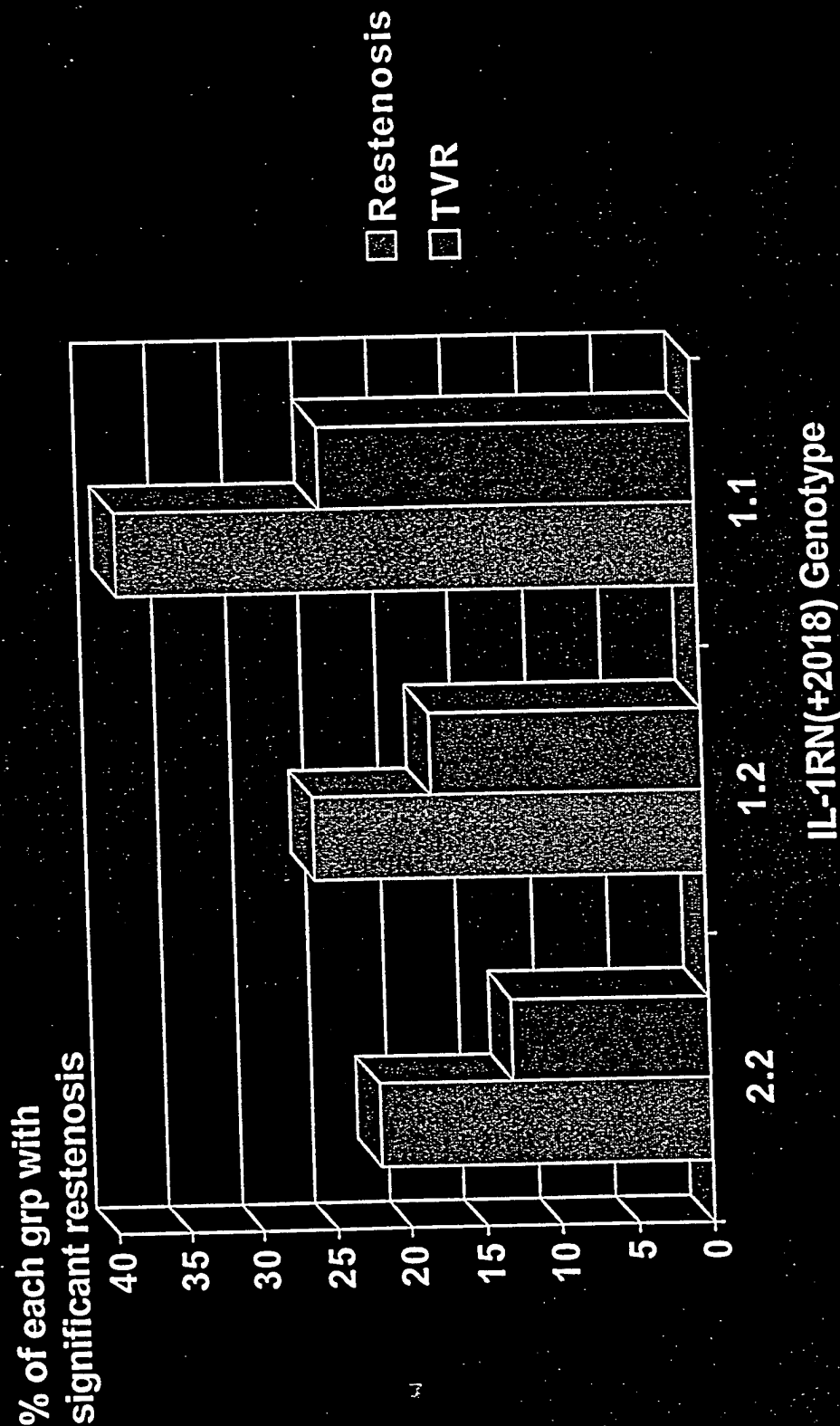


Figure 8

A Different Pattern is Associated with Risk for Restenosis: Preliminary data

Kastrati, Schomig, et al 1999



004250 4587560

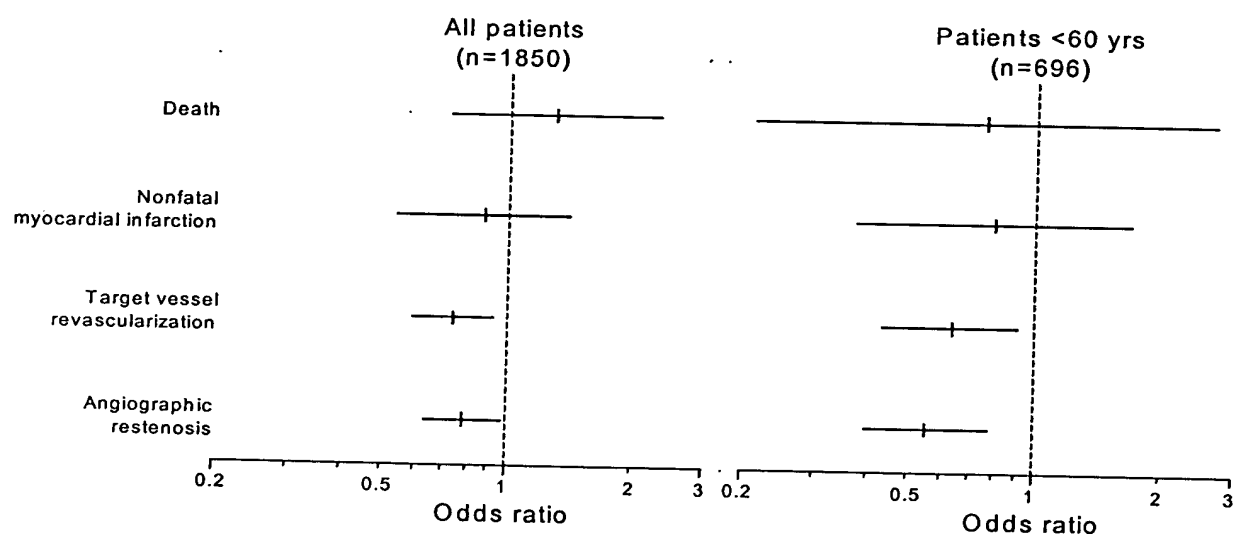


Figure 9

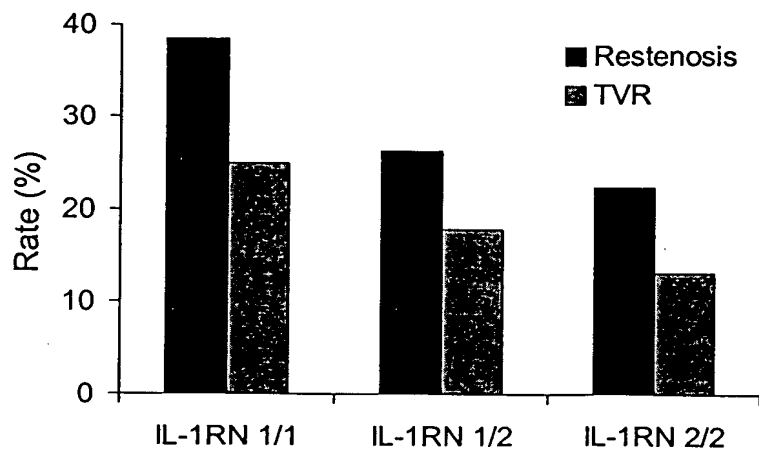


Figure 10